=> s (IL-1ra or interleukin-1 receptor antagonist#)

LI 5227 (IL-1RA OR INTERLEUKIN-1 RECEPTOR ANTAGONIST#)

=> s (IL-1ra-R or interleukin-1 receptor antagonist related)

L2 14 (IL-1RA-R OR INTERLEUKIN-1 RECEPTOR ANTAGONIST RELATED)

=> d 12 1-12 bib ab

L2 ANSWER 1 OF 14 MEDLINE

AN 2001138840 MEDLINE

DN 21030891 PubMed ID: 11192058

TI Physical activity and plasma interleukin-6 in humans--effect of intensity

of exercise.

AU Ostrowski K; Schjerling P; Pedersen B K

CS The Copenhagen Muscle Research Centre, Rigshospitalet Afs 7652, Denmark.

SO Eur J Appl Physiol, (2000 Dec) 83 (6) 512-5.

Journal code: 100954790. ISSN: 1439-6319.

CY Germany: Germany, Federal Republic of

DT Journal; Article; (JOURNAL ARTICLE)

LA English

FS Priority Journals

EM 200103

ED Entered STN: 20010404 Last Updated on STN: 20010404

Entered Medline: 20010308

AB The present study included data from three marathon races to investigate

the hypothesis that a relationship exists between running intensity and

elevated concentrations of interleukin (IL)-6 in plasma. The study included a total of 53 subjects whose mean age was 30.6 [95% confidence

interval (CI) 1.4] years, mean body mass 77.7 (95% CI 2.0) kg, mean

maximal oxygen uptake (VO2max) 59.3 (95% Cl 1.4) ml x min(-1) x kg(-1),

and who had participated in the Copenhagen Marathons of 1996, 1997 or

1998, achieving a mean running time of 206 (95% CI 7) min. Running

intensity was calculated as running speed divided by VO2 max. The

concentration of IL-6 in plasma peaked immediately after the run. There

was a negative correlation between peak IL-6 concentration and running

time (r = -0.30, P<0.05) and a positive correlation between peak IL-6

concentration and running intensity (r = 0.32, P<0.05). The IL-1 receptor

antagonist (IL-1ra) plasma concentration peaked 1.5 h after the run and

there was a positive correlation between the peak plasma concentrations of

IL-6 and ***IL*** - ***1ra*** (***r*** = 0.39, P<0.01). Creatine

kinase (CK) plasma concentration peaked on the 1st day after the run, but

no association was found between peak concentrations of IL-6 and CK, In

conclusion, the results confirmed the hypothesized association between

plasma IL-6 concentration and running intensity, but did not confirm the

previous finding of a connection between IL-6 plasma concentration and

muscle damage.

L2 ANSWER 2 OF 14 MEDLINE

AN 97225342 MEDLINE

DN 97225342 PubMed ID: 9071715

TI Lipopolysaccharide-binding protein and

bactericidal/permeability-

increasing factor during hemodialysis: clinical determinants and role of

different membranes.

AU Sundaram S; King A J; Pereira B J

CS Division of Nephrology, New England Medical Center, Boston, Massachusetts

02111, USA.

NC DK 45609 (NIDDK)

SO JOURNAL OF THE AMERICAN SOCIETY OF

NEPHROLOGY, (1997 Mar) 8 (3) 463-70.

Journal code: 9013836. ISSN: 1046-6673.

CY United States

DT Journal; Article; (JOURNAL ARTICLE)

LA English

FS Priority Journals

EM 199706

ED Entered STN: 19970620

Last Updated on STN: 19970620

Entered Medline: 19970611

AB The host response to the presence of lipopolysaccharide (LPS) is complex

and varied. Two closely related endogenous serum proteins,

LPS-binding

protein (LBP) and bactericidal/permeability-increasing factor (BPI),

regulate delivery of LPS to CD14 antigen on effector cell surfaces and

modulate the host response to LPS. In the study presented here, plasma

levels of LBP and BPI were measured, predialysis, 15 min into dialysis and

postdialysis in patients dialyzed with cellulose,

cellulose-tri-acetate

(CTA), and polysulfone dialyzers. Further, the association between LBP

levels and BPI release during hemodialysis and clinical and laboratory

characteristics of patients, complement activation represented by plasma

C3a levels, and monocyte cytokine production represented by interleukin-1

receptor antagonist (IL-1Ra) synthesis was also studied.

Predialysis

plasma levels of LBP were 14,459 +/- 544, 13,889 +/- 1362 and 12,622 +/-

6305 ng/mL, respectively, with cellulose, CTA, and polysulfone dialyzers,

and postdialysis levels were 17,834 + /-861, 20,979 + /-8485 and 18,177

+/- 1656 ng/mL, respectively. Postdialysis plasma levels of LBP

consistently higher than predialysis levels with all three dialyzers (P <

0.05). However, plasma LBP levels were not significantly different between

the three dialyzers either predialysis (P = 0.28) or postdialysis (P = 0.28)

2.8). There were no significant differences in predialysis BPI levels ${\bf P}_{\bf P}$

between the three dialyzers (P = 0.21). BPI levels at 15 min of

dialysis

with CTA (10.91 +/- 3.65 ng/mL) and polysulfone (10.73 +/- 2.24 ng/mL)

dialyzers were significantly greater (P < 0.05) than that observed

cellulose (5.49 +/- 0.66 ng/mL). Similarly, postdialysis levels with CTA

and polysulfone were significantly greater (P < 0.05) than that observed

with cellulose dialyzers. The percentage change in BPI levels

predialysis and 15 min was 1341 +/- 243%, 2935 +/- 1033%, and 3790 +/-

1151% for cellulose, CTA, and polysulfone dialyzers, respectively. The

changes in BPI levels from predialysis to 15 min and between preand

postdialysis samples were statistically significant for all three dialyzers (P < 0.05). Postdialysis LBP:BPI ratios were 50 +/- 6%, 18 +/-

4%, and 22 +/- 6% of predialysis ratios for cellulose, CTA, and polysulfone dialyzers, respectively. These changes were statistically

significant (P < 0.05) for all three dialyzers. There was no significant

correlation between baseline clinical or laboratory characteristics and

predialysis LBP levels. Similarly, the correlation between BPI levels at

15 min of dialysis with the clinical and laboratory characteristics was

also poor, with the exception of serum albumin (r = 0.43, P = 0.008). The

correlation between BPI levels at 15 min of dialysis with plasma LBP

levels (r = -0.29; P = 0.08), plasma C3a levels (r = -0.1; P = 0.55),

peripheral blood mononuclear cells (PBMC) content of ***IL***

1Ra (***r*** = 0.01; P = 0.94), and IL-1Ra production by

unstimulated (r = 0.13; P = 0.45), and endotoxin-stimulated PBMC (r =

0.32; P = 0.06) was not statistically significant. The results of this study demonstrate that dialysis with cellulose, CTA, and polysulfone

dialyzers results in a significant increase in LBP and BPI levels.

release is probably mediated by non-complement factors and may

to the nutritional status of the patient. The release of BPI during HD and

consequent lowering of the LBP:BPI ratio could potentially afford some

protection against endotoxin in the dialysate.

L2 ANSWER 3 OF 14 MEDLINE

AN 96416422 MEDLINE

DN 96416422 PubMed ID: 8928570

TI [Practical significance of cytokine determination in joint fluid in patients with arthroses or rheumatoid arthritis].

Praktische Bedeutung der Zytokinbestimmung im Gelenkpunktat von Patienten

mit Arthrosen oder rheumatischen Arthritiden.

AU Neidel J; Schulze M; Sova L; Lindschau J

CS Abt. fur Orthopadie, Rheumaklinik Bad Bramstedt, Medizinische Hochschule

Hannover.

SO ZEITSCHRIFT FUR ORTHOPADIE UND IHRE GRENZGEBIETE, (1996 Jul-Aug) 134 (4)

381-5.

Journal code: 1256465. ISSN: 0044-3220. CY GERMANY: Germany, Federal Republic of

DT Journal; Article; (JOURNAL ARTICLE)

LA German

FS Priority Journals

EM 199611

ED Entered STN: 19961219 Last Updated on STN: 20000303

Entered Medline: 19961114

AB OBJECTIVE: To determine whether the activity of cartilage-degrading

enzymes in the synovial fluid (SF) of patients with rheumatoid arthritis

and other joint diseases is correlated with the concentration of cytokines

in the SF. METHODS: Cytokines and cartilage-degrading enzymes were

determined in the SF of 97 patients with various disorders involving the

knee joints (rheumatoid arthritis (RA) n 44; osteoarthritis (OA) n 35;

meniscal trauma (Men) n 10; reactive arthritides (ReA) n 8). In

samples we measured the concentrations of interleukin-1 alpha and beta,

IL-1-receptor antagonist (IL-1ra), IL-6, IL-8, tumor necrosis factor alpha

(TNF alpha; all by ELISA), collagenase-activity and caseinase-activity (by

substrate assays). RESULTS: With the exception of IL-1 alpha and IL-6,

cytokine-concentrations were significantly higher in RA than in OA

SF-samples (p < 0.05; ANOVA on ranks). IL-1ra, IL-6, and IL-1 beta were

correlated best with the collagenase-activity in the SF (r = 0.63; 0.57;

0.55; Spearman's rank correlation), while IL-1 beta (r = 0.53) and ***IL*** - ***1ra*** (***r*** = 0.52) were best correlated with

the case inase-activity in the samples. The SF-concentration of IL-1 ra was $\label{eq:ll-1}$

well correlated with the levels of IL-6, IL-1 beta, II-8, and TNF alpha (r $\,$

from 0.73 to 0.66; all p < 0.005), but not with IL1 alpha. The molar ratio of IL-1 to IL-1ra in the SF was neither correlated with the activity

of collagenase nor caseinase. IL-1 beta and IL-1ra in the SF were

positively correlated with the erythrocyte sedimentation rate (ESR).

CONCLUSIONS: The determination of IL-1 beta and IL-1ra in the SF of patients with

disorders as examined in this study seems to allow to a certain

extent a prediction of the collagenase- and caseinase-activity contained in

diseased joint. We would favor.

L2 ANSWER 4 OF 14 MEDLINE

AN 96188960 MEDLINE

DN 96188960 PubMed ID: 8608647

TI Significance of IL-1 beta and IL-1 receptor antagonist (IL-1Ra) in

bronchoalveolar lavage fluid (BALF) in patients with diffuse panbronchiolitis (DPB).

AU Kadota J; Matsubara Y; Ishimatsu Y; Ashida M; Abe K; Shirai R; Iida K;

Kawakami K; Taniguchi H; Fujii T; Kaseda M; Kawamoto S; Kohno S

CS Second Department of Internal Medicine, Nagasaki University School of

Medicine, Japan.

SO CLINICAL AND EXPERIMENTAL IMMUNOLOGY, (1996 Mar) 103 (3) 461-6.

Journal code: 0057202. ISSN: 0009-9104.

CY ENGLAND: United Kingdom

DT Journal; Article; (JOURNAL ARTICLE)

LA English

FS Priority Journals

EM 199605

ED Entered STN: 19960605 Last Updated on STN: 19960605 Entered Medline: 19960528

AB We evaluated the effect of erythromycin therapy on pulmonary function

tests and the airway inflammatory response of patients with DPB. The

number of neutrophils in BALF obtained from DPB patients was significantly

higher than that of healthy volunteers. Treatment with erythromycin (600

mg/day for 12.9+/-9.5 months (mean +/- s.d.)) significantly reduced the

total number of cells and neutrophils in the airway, and significantly

improved pulmonary function tests. The levels of IL-1 beta and IL-8 were

significantly higher in DPB compared with healthy volunteers (P<0.05,

P<0.05, respectively). IL-1Ra in patients is considered to have a weak

inhibitory activity for IL-1beta, with approximately five-fold concentration of IL-1beta compared with that in healthy volunteers

(approx. nine-fold concentration of IL-1beta). Erythromycin therapy

significantly reduced these cytokines to levels comparable to those of

healthy volunteers, and produced a trend toward reduction in the level of

IL-1Ra in BALF. The level of IL-1beta correlated significantly with the

concentration of neutrophils in BALF (r=0.72, P<0.01), as well as with the

level of ****IL*** - ****1Ra*** (***r*** =0.688, P<0.05) and IL-8

(r=0.653, P<0.05). A nearly significant or significant correlation was

observed between the concentration of neutrophils and levels of IL-1Ra or

IL-8 in BALF (r=0.526, P=0.053 or r=0.776, P<0.01, respectively). There

was also a significant relationship between FEV(1) and the concentration

of neutrophils in BALF (r=0.524, P<0.05). Our results suggest that the

relative amounts of IL-1beta and IL-1Ra or IL-8 may contribute, at least

in part, to the neutrophil-mediated chronic airway inflammation in patients with chronic airway disease, and long-term erythromycin terapy

may down-regulate the vigorous cycle between the cytokine network and

neutrophil accumulation, with resultant reduction of neutrophil-mediated

inflammatory response.

L2 ANSWER 5 OF 14 MEDLINE

AN 95189896 MEDLINE

DN 95189896 PubMed ID: 7883859

TI Soluble cytokine receptors and the low 3,5,3'-triiodothyronine syndrome in

patients with nonthyroidal disease.

AU Boelen A; Platvoet-Ter Schiphorst M C; Wiersinga W M CS Department of Endocrinology, University of Amsterdam, The Netherlands.

SO JOURNAL OF CLINICAL ENDOCRINOLOGY AND

METABOLISM, (1995 Mar) 80 (3) 971-6.

Journal code: 0375362. ISSN: 0021-972X.

CY United States

DT Journal; Article; (JOURNAL ARTICLE)

LA English

FS Abridged Index Medicus Journals; Priority Journals

EM 199504

ED Entered STN: 19950425 Last Updated on STN: 19950425 Entered Medline: 19950411

AB Cytokines have been implicated in the pathogenesis of the low T3 syndrome

during illness. This is supported by our recent observation of a strong

negative relationship between serum T3 and serum interleukin-6 (IL-6) in

nonthyroidal illness (NTI). In the last few years, soluble cytokine receptors and cytokine receptor antagonists have been discovered in human

serum. These proteins have the potential to further regulate cytokine

activity. Therefore, we now studied the association between serum T3 and

serum levels of soluble tumor necrosis factor-alpha (sTNF alpha R p55 and

sTNF alpha R p75), soluble interleukin-2 receptor (sIL-2R), and the

interleukin-1 receptor antagonist (IL-1RA) in 100 consecutive hospital

admissions with a wide variety of nonthyroidal diseases. Patients were

divided into group A (T3, > or = 1.30 nmol/L; T4, > or = 75 nmol/L; n =

41), group B (T3, < 1.30 nmol/L; T4, > or = 75 nmol/L; n = 46), and group

C (T3, < 1.30 nmol/L; T4, < 75 nmol/L; n = 13). Serum sTNF alpha R p55, sTNF alpha R p75, sIL-2R, and IL-1RA were lower in group A

than in groups
B and C [median values; sTNF alpha R p55, 1.25, 2.25, and 3.55

ng/mL (P < 0.001); sTNF alpha R p75, 2.02, 4.56, and 7.00 ng/mL (P <

0.001); sIL-2R, 184, 259, and 272 U/mL (P = 0.0004), respectively]. Serum IL-1RA levels

were not different in the three groups (median values, 122, 193, and 258 $\,$

and 258 pg/mL, respectively). Taking all patients together, a significant

negative relation was found among serum T3 and sTNF alpha p55 (r =

-0.59; P < 0.0001), sTNF alpha R p75 (r = -0.55; P < 0.0001), sIL-2R (r =

-0.54; P < 0.0001), ***IL*** - ***1RA*** (***r*** = -0.38; P = 0.001), and

IL-6 (r = -0.56; P < 0.0001). A remarkable high correlation (r = -0.70; P

< 0.0001) was found between serum T3 and a newly designed total score

based on the summation of serum levels of IL-6 and the four soluble

cytokine receptor proteins. IL-6 and the four cytokine receptor proteins

were all significantly related to each other. Stepwise multiple

regression underlying leukocyte activation in this disorder. The increased cytokine indicated IL-6 and sTNF alpha R p75 as independent concentration may also be responsible for the endothelial adhesion determinants of T3 [serum T3 = 2.09-0.32ln (sTNF alpha R p75) -0.15ln (IL-6); r = that 0.701. The accompanies preeclampsia. variability in serum T3 was accounted for 35% by changes in In L2 ANSWER 7 OF 14 CAPLUS COPYRIGHT 2002 ACS (sTNF alpha AN 2001:435130 CAPLUS R p75) and 14% by changes in ln (IL-6) (ABSTRACT TRUNCATED AT 400 WORDS) DN 135:41824 TI DNA encoding human and murine ***interleukin*** -L2 ANSWER 6 OF 14 MEDLINE AN 95060548 MEDLINE DN 95060548 PubMed ID: 7526306 IN Saris, Christian M.; Giles, Jennifer; Mu, Sharon X.; Xia, Min; TI Increased concentrations of cytokines interleukin-6 and interleukin-1 Michael Brian; Craveiro, Roger receptor antagonist in plasma of women with preeclampsia: a PA Amgen, Inc., USA mechanism for SO PCT Int. Appl., 190 pp. endothelial dysfunction?. CODEN: PIXXD2 AU Greer I A; Lyall F; Perera T; Boswell F: Macara L M DT Patent CS Department of Obstetrics and Gynecology, Royal Infirmary, LA English FAN.CNT 1 Glasgow. Scotland, United Kingdom. PATENT NO. KIND DATE APPLICATION NO. SO OBSTETRICS AND GYNECOLOGY, (1994 Dec) 84 (6) DATE 937-40. Journal code: 0401101. ISSN: 0029-7844. PI WO 2001042304 A1 20010614 WO 2000-US32940 CY United States DT Journal; Article; (JOURNAL ARTICLE) W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, LA English BZ, CA, CH, CN, FS Abridged Index Medicus Journals; Priority Journals CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, EM 199412 GH, GM, HR, ED Entered STN: 19950110 HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, Last Updated on STN: 19960129 LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, Entered Medline: 19941213 AB OBJECTIVE: To determine if plasma concentrations of defined PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, cytokines are increased in women with preeclampsia, and to correlate any UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM increases with the elevated concentrations of the vascular cell adhesion molecule RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, (VCAM)-1. METHODS: Twenty primigravidas with AT, BE, CH, CY, preeclampsia were compared to DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, 20 healthy primigravidas. Plasma levels of cytokines, tumor TR, BF, necrosis BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, factor-alpha (TNF alpha), interleukin (IL)-6, IL-8, IL-1 beta, IL-1 TG PRAI US 1999-170191P P 19991210 receptor antagonist (IL-1ra), granulocyte macrophage-colony-stimulating US 2000-188053P P 20000309 factor (GM-CSF), and VCAM-1, were measured by US 2000-194521P P 20000404 enzyme-linked immunosorbent US 2000-195910P P 20000410 AB The present invention provides nucleic acid mols. encoding assay. RESULTS: Concentrations of IL-6 and IL-1ra were significantly ***Interleukin*** - ***1*** ***Receptor*** higher (P < .01) in preeclamptic women (2.56 and 251.85 pg/mL, ***Antagonist*** respectively) compared to normal pregnant patients (2.06 and ***Related*** (***IL*** - ***1ra*** - ***R***) 142.00 pg/mL, respectively). There were no significant changes in concentrations The cDNAs encoding human and murine ***IL*** - ***1ra*** ***R*** alpha, IL-8, GM-CSF, and IL-1 beta in preeclamptic patients (14.09, 50.52, were cloned and the expression in several human tissues were 125.8, and 2.08 pg/mL, respectively) compared to normal patients examd. by either RT-PCR or in situ hybridization. ***IL*** - ***1ra*** (11.96 44.46, 121.3, and 2.01 pg/mL, respectively). Serum ***R*** was expressed in E. coli and mammalian cell and anticoncentrations of VCAM-1 were increased in women with preeclampsia ***IL*** (preeclamptic group 841.9 - ***1ra*** - ***R*** antibody was produced. The biol. +/-49.7 ng/mL, control group 560.2 +/-47.9 ng/mL; t = 3.673, P activity of ***IL*** - ***1ra*** - ***R*** was assessed in transgenic Interleukin-6 and IL-1ra concentrations correlated with VCAM-1 mice. The concentrations (IL-6: r = 0.539, z = 2.9, P < .005; ***IL*** invention also provides selective binding agents, vectors, host ***1ra***: ***r*** = 0.451, z = 2.428, P < .02).

methods for producing ***IL*** - ***1ra*** - ***R***

The invention further provides pharmaceutical compns. and

methods for the

CONCLUSIONS:

endothelial damage

Increased cytokine concentrations may contribute to the

that occurs with preeclampsia and may explain the mechanism

diagnosis, treatment, amelioration, and/or prevention of diseases, disorders, and conditions assocd. with ***IL*** - ***lra*** - ***R*** polypeptides.

RE.CNT 6 THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD

ALL CITATIONS AVAILABLE IN THE RE FORMAT

L2 ANSWER 8 OF 14 CAPLUS COPYRIGHT 2002 ACS

AN 1996:259086 CAPLUS

DN 124:331889

TI Significance of IL-1.beta. and IL-1 receptor antagonist (IL-1Ra) in

bronchoalveolar lavage fluid (BALF) in patients with diffuse panbronchiolitis (DPB)

AU Kadota, J.; Matsubara, Y.; Ishimatsu, Y.; Ashida, M.; Abe, K.; Shirai, R.;

Iida, K.; Kawakami, K.; Taniguchi, H.; et al.

CS School Medicine, Nagasaki University, Nagasaki, 852, Japan

SO Clin. Exp. Immunol. (1996), 103(3), 461-6 CODEN: CEXIAL; ISSN: 0009-9104

DT Journal

LA English

AB We evaluated the effect of erythromycin therapy on pulmonary function

tests and the airway inflammatory response of patients with DPB. The no.

of neutrophils in BALF obtained from DPB patients was significantly higher

than that of healthy volunteers. Treatment with erythromycin (600 mg/day

for 12.sum.9.+-.9.sum.5 mo (mean .+-. s.d.)) significantly reduced the

total no. of cells and neutrophils in the airway, and significantly improved pulmonary function tests. The levels of IL-1.beta. and IL-8 were

significantly higher in DPB compared with healthy volunteers (P

0.sum.05, P < 0.sum.05, resp.). IL-1Ra in patients is considered to have

a weak inhibitory activity for IL-1.beta., with approx. five-fold conen.

of IL-1.beta. compared with that in healthy volunteers (approx. nine-fold

concn. of IL-1.beta.). Erythromycin therapy significantly reduced

cytokines to levels comparable to those of healthy volunteers, and produced a trend toward redn. in the level of IL-1Ra in BALF. The level

of IL-1.beta. correlated significantly with the concn. of neutrophils in

BALF (r = 0.72, P < 0.01), as well as with the level of ***IL***

1Ra (***r*** = 0.688, P < 0.05) and IL-8 (r = 0.653,

0.05). A nearly significant or significant correlation was obsd. between

the concn. of neutrophils and levels of IL-1Ra or IL-8 in BALF (r = 0.526.

P = 0.053 or r = 0.776, P < 0.01, resp.). There was also a significant

relation between FEV1 and the concn. of neutrophils in BALF (r = 0.524, P

 $\!<\!0.05).$ Our results suggest that the relative amts. of IL-1.beta, and

IL-1Ra or IL-8 may contribute, at least in part, to the neutrophil-mediated chronic airway inflammation in patients with chronic

airway disease, and long-term erythromycin therapy may

vigorous cycle between the cytokine network and neutrophil accumulation,

with resultant redn. of neutrophil-mediated inflammatory response.

L2 ANSWER 9 OF 14 CAPLUS COPYRIGHT 2002 ACS AN 1995:437486 CAPLUS

TI Soluble cytokine receptors and the low 3,5,3'-triiodothyronine syndrome in

patients with nonthyroidal disease

AU Boelen, A.; Schiphorst, M. C. Platvoet-ter; Wiersinga, W. M.

CS Department of Endocrinology, Univ. of Amsterdam, Amsterdam, Neth.

SO J. Clin. Endocrinol. Metab. (1995), 80(3), 971-6 CODEN: JCEMAZ; ISSN: 0021-972X

DT Journal

LA English

AB Cytokines have been implicated in the pathogenesis of the low T3 syndrome

during illness. This is supported by our recent observation of a strong

neg. relationship between serum Tc and serum interleukin-6 (IL-6) in

nonthyroidal illness (NTI). In the last few years, sol. cytokine receptors and cytokine receptor antagonists have been discovered in human

serum. These proteins have the potential to further regulate cytokine

activity. Therefore, we now studied the assocn. between serum T3

serum levels of sol. tumor necrosis factor-.alpha. receptors (sTNF.alpha.R

p55 and sTNF.alpha.R p75), solbule interleukin-2 receptor (sIL-2R), and

the interleukin-1 receptor antagonist (IL-1RA) in 100 consecutive hospital

admissions with a wide variety of nonthyroidal diseases. Patients were

divided into group A (T3, .gtoreq.1.30 nmol/L; T4, .gtoreq.75 nmol/L; n =

41), group B (T3, <1.30 nmol/L; T4, .gtoreq.75 nmol/L; n = 46), and group

C (T3, <1.30 nmol/L; T4, <75 nmol/L; n = 13). Serum sTNF.alpha.R p55,

sTNF.alpha.R p75, sIL-2R, and IL-1RA were lower in group A than in groups

B and C [median values: sTNF.alpha.R p55, 1.26, 2.25, and 3.55 ng/mL (P <

0.001); sTNF.alpha.R p75, 2.02, 4.56, and 7.00 ng/mL (P < 0.001); sIL-2R,

184, 259, and 272 U/mL (P = 0.0004), resp.]. Serum IL-1RA levels were not different in the three groups (median values 122, 193, and 258

different in the three groups (median values, 122, 193, and 258 pg/mL,

resp.). Taking all patients together, a significant neg. relation was found among serum T3 and sTNF.alpha. p55 (r = -0.59; P < 0.0001),

sTNR.alpha.R p75 (r = -0.55; P < 0.0001), sIL-2R (r = -0.54; P < 0.0001),

IL - ***1RA*** (***r*** = -0.38; P = 0.001), and IL-6 (r =

-0.56; P < 0.0001). A remarkable high correlation (r = -0.70; P < 0.0001)

was found between serum T3 and a newly designed total score based on the

summation of serum levels of IL-6 and the four sol. cytokine

proteins. IL-6 and the four cytokine receptor proteins were all significantly related to each other. Stepwise multiple regression indicated IL-6 and sTNF.alpha.R p75 as independent determinants of T3

[serum T3 = 2.09 - 0.32ln (sTNF.alpha.R p75) - 0.15ln (IL-6); r = 0.701.

The variability in serum T3 was accounted for 35% by changes in ln

(sTNF.alpha.R p75) and 14% by changes in ln (IL-6). In conclusion, 1)

serum T3 is neg. related to serum sTNF.alpha.R p55, sTNF.alpha.R p75,

sIL-2R, IL-1RA, and IL-6 in patients; and 2) sTNF.alpha.R p75 and IL-6 are

independent determinants of serum T3 in NTI, accounting for 35% and 14%,

resp., of the variability in T3. The results suggest that the sick euthyroid syndrome is part of the acute phase response during

generated by activation of the cytokine network.

L2 ANSWER 10 OF 14 USPATFULL

AN 2002:5759 USPATFULL

TI Interleukin-1 receptor antagonist and recombinant production thereof

Ford, John, San Mateo, CA, United States Pace, Ann, Scotts Valley, CA, United States

Hyseq, Inc., Sunnyvale, CA, United States (U.S. corporation)

US 6337072 PΙ

B1 20020108

19990707 (9) US 1999-348942 ΑI

RLI Continuation-in-part of Ser. No. US 1999-287210, filed on 5 Apr 1999.

now abandoned Continuation-in-part of Ser. No. US 1999-251370, filed on

17 Feb 1999, now abandoned Continuation-in-part of Ser. No.

1999-229591, filed on 13 Jan 1999, now abandoned Continuation-in-part of

Ser. No. US 1998-127698, filed on 31 Jul 1998, now abandoned Continuation of Ser. No. US 1998-99818, filed on 19 Jun 1998,

abandoned Continuation of Ser. No. US 1998-82364, filed on 20 May 1998,

now abandoned Continuation-in-part of Ser. No. US 1998-79909, filed on

15 May 1998, now abandoned Continuation-in-part of Ser. No.

1998-55010, filed on 3 Apr 1998, now abandoned

PRAI WO 1999-US4291 19990405

DT Utility

GRANTED

EXNAM Primary Examiner: Spector, Lorraine

LREP Marshall, O'Toole, Gerstein, Murray & Borun

CLMN Number of Claims: 37

ECL Exemplary Claim: 1,15

DRWN 4 Drawing Figure(s); 4 Drawing Page(s)

LN.CNT 5025

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The present invention provides novel nucleic acids, the novel polypeptide sequences encoded by these nucleic acids and uses thereof.

These novel polynucleotide and polypeptide sequences were determined to

be a novel Interleukin-1 Receptor Antagonist.

L2 ANSWER 11 OF 14 USPATFULL

AN 2001:163320 USPATFULL

TI Anti-interleukin-1 receptor antagonist antibodies and uses thereof

Ford, John, San Mateo, CA, United States IN Pace, Ann, Scotts Valley, CA, United States

PA Hyseq, Inc., Sunnyvale, CA, United States (U.S. corporation)

B1 20010925 US 6294655

US 1999-417455 19991013 (9) ΑĪ

RLI Continuation-in-part of Ser. No. US 1999-348942, filed on 7 Jul 1999

Continuation of Ser. No. US 1999-287210, filed on 5 Apr 1999,

abandoned Continuation-in-part of Ser. No. US 1999-251370, filed on 17

Feb 1999, now abandoned Continuation-in-part of Ser. No. US 1998-127698,

filed on 31 Jul 1998, now abandoned Continuation-in-part of Ser. No. US

1999-229591, filed on 13 Jan 1999, now abandoned Continuation of Ser.

No. US 1998-99818, filed on 19 Jun 1998, now abandoned . said Ser. No.

US 127698 Continuation-in-part of Ser. No. US 1998-82364, filed on 20

May 1998, now abandoned, said Ser. No. US 99818 Continuation-in-part of

Ser. No. US 1998-82364, filed on 20 May 1998, now abandoned

Continuation-in-part of Ser. No. US 1998-79909, filed on 15 May 1998,

now abandoned Continuation-in-part of Ser. No. US 1998-55010, filed on 3

Apr 1998, now abandoned

. Utility DT

GRANTED

EXNAM Primary Examiner: Spector, Lorraine

LREP Marshall, O'Toole Gerstein, Murray & Borun

CLMN Number of Claims: 14

ECL Exemplary Claim: 1

DRWN 15 Drawing Figure(s); 14 Drawing Page(s)

LN.CNT 4656

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The present invention provides novel nucleic acids, the novel polypeptide sequences encoded by these nucleic acids and uses thereof.

These novel polynucleotide and polypeptide sequences were determined to

be a novel Interleukin-1 Receptor Antagonist. Also provided are antibodies which bind the antagonist, methods of detecting the antagonist, and kits containing the antibodies.

L2 ANSWER 12 OF 14 USPATFULL

AN 1999:132765 USPATFULL

TI Method of treatment of osteoarthritis with interleuken-1 receptor

antagonist

Pelletier, Jean-Pierre, St-Lambert, Canada Martel-Pelletier, Johanne, St-Lambert, Canada

Arthro Lab Inc., Sherbrooke, Canada (non-U.S. corporation) 19991026

US 5972880 PΙ ΑI

US 1996-612433 19960307 (8)

DT Utility

FS Granted

EXNAM Primary Examiner: Mertz, Prema

LREP ROBIC

CLMN Number of Claims: 3

ECL Exemplary Claim: 1

DRWN 2 Drawing Figure(s); 2 Drawing Page(s)

LN.CNT 745

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

A method and a composition for the preventative treatment of osteoarthritis comprising the periodic administration to a mammal

suffering of this disease of a composition comprising an amount of Human

recombinant Interleukin-1 receptor antagonist effective for reducing the

progression of lesions and cartilage degradation.